

AMENDMENTS TO THE CLAIMS

This listing of claims will replace all prior versions, and listings, of claims in the application:

5 LISTING OF CLAIMS:

1. - 58. (Canceled)

59. (currently amended) A method of determining polymorphic sites or sub-haplotypes for a locus that correlate with a clinical response or an outcome of interest, comprising:

- 10 (a) providing a haplotype information, for each allele of the locus for each subject in a cohort of subjects and clinical response or outcome data (clinical outcome values) an outcome value for the outcome of interest for each subject in the from a cohort of subjects, the locus comprising at least two polymorphic sites;
- 15 (b) statistically analyzing each individual SNP polymorphism in the haplotype-haplotypes for the degree to which it correlates with the outcome of interest clinical outcome values, and generating calculating a numerical measure of the degree of correlation;
- 20 (c) saving for further processing these each individual SNPs polymorphism whose numerical measure of the degree of correlation with the clinical outcome values exceeds-meets a first cut-off value criterion and its numerical measure of the degree of correlation;
- 25 (d) generating all possible pair-wise combinations of the saved SNPs individual polymorphisms so as to provide a set of n -site sub-haplotypes where $n = 2$;
- (e) statistically analyzing each newly generated n -site sub-haplotype for the degree to which it correlates with the clinical outcome values outcome of interest and calculating a the numerical measure of the degree of correlation;
- 30 (f) saving for further processing these each n -site sub-haplotypes haplotype whose numerical measure of the degree of correlation with the clinical outcome values exceeds-meets the first cut-off value criterion and its numerical measure of the degree of correlation;
- (g) generating all possible pair-wise combinations among and between the saved SNPs

individual polymorphisms and saved sub-haplotypes, to produce new ~~subhaplotypes~~
sub-haplotypes with increased values of n ; and

- (h) repeating steps (e) through (g) until either (i) no new sub-haplotypes can be generated,
or (ii) no ~~further-new~~ sub-haplotypes having n less than a pre-selected limit can be
generated.

60. (currently amended) The method of claim 59, further comprising the step of displaying
~~these the saved SNPs~~ individual polymorphisms and sub-haplotypes whose numerical
measure of the degree of correlation ~~with the clinical outcome value exceeds~~ meets a
second cut-off ~~value~~ criterion, wherein the second cut-off ~~value~~ criterion is ~~greater~~ more
stringent than the first cut-off ~~value~~ criterion.

61. (currently amended) The method of claim 59, wherein the numerical measure of the
degree of correlation is ~~replaced by the a p-value for the correlation~~, and SNPs and sub-
haplotypes are saved if the first cut-off criterion is that the p-value is less than or equal to a
first cut-off value.

62. (currently amended) The method of claim 61, further comprising the step of displaying
~~these the saved SNPs~~ individual polymorphisms and sub-haplotypes whose p-value for the
correlation ~~with the clinical outcome value is less than~~ or equal to a second cut-off value,
wherein the second cut-off value is less than the first ~~selected~~ cut-off value.

63. (currently amended) The method of any one of claims 59-62, further comprising the step
of ~~excluding from further processing~~ eliminating a complex redundant sub-haplotype
~~subhaplotypes which are constructed-generated~~ from smaller saved sub-haplotypes, ~~where~~
wherein the numerical measure of the degree of correlation of each of the smaller saved
sub-haplotypes each have correlation values that are is at least as significant as that of the
complex redundant sub-haplotype.

64. (currently amended) A method of determining polymorphic sites or sub-haplotypes for a
locus that correlate with a clinical response ~~or an~~ outcome of interest, comprising:

- (a) providing a single gene haplotype for each allele of the locus for each subject in a cohort of subjects information for one or more genes, and clinical response or outcome data, from a cohort of subjects an outcome value for the outcome of interest for each subject in the cohort, the locus comprising at least two polymorphic sites;
- (b) statistically analyzing each single gene haplotype for the degree to which it correlates with the clinical response or outcome of interest, and calculating a numerical measure of the degree of correlation;
- (c) saving for further processing those haplotypes each haplotype whose numerical measure of the degree of correlation with the clinical response or outcome of interest meets exceeds a first selected value cut-off criterion and its numerical measure of the degree of correlation;
- (d) for from each haplotype composed of m polymorphic sites, generating all possible sub-haplotypes having a single site masked, so as to provide a set of sub-haplotypes having $(m-n)$ sites, where $n = 1$;
- (e) statistically analyzing each newly generated sub-haplotype for the degree to which it correlates with the clinical response or outcome of interest, and calculating a the numerical measure of the degree of correlation;
- (f) saving for further processing those each sub-haplotypes haplotype whose numerical measure of the degree of correlation with the clinical response or outcome of interest exceeds meets the first selected value cut-off criterion and its numerical measure of the degree of correlation;
- (g) from the saved sub-haplotypes, generating all possible sub-haplotypes having one additional site masked; and
- (h) repeating steps (e) through (g) until either (i) no new sub-haplotypes have a degree of correlation which exceeds the first selected value, or (ii) no further sub-haplotypes having more unmasked sites than a pre-selected limit can be generated.

65. (currently amended) The method of claim 64, further comprising the step of displaying those saved haplotypes and sub-haplotypes whose numerical measure of the degree of

~~correlation with the clinical response or outcome of interest exceeds~~ meets a second
~~selected value cut-off criterion, wherein the second selected value cut-off criterion is greater~~
~~more stringent than the first selected value cut-off criterion.~~

- 5 66. (currently amended) The method of claim 64, wherein the numerical measure of the
degree of correlation is replaced by ~~the~~ a p-value for the correlation, and the first cut-off
criterion is that sub-haplotypes are saved if the p-value is less than or equal to a first
selected value.
- 10 67. (currently amended) The method of claim 66, further comprising the step of displaying
those saved haplotypes and sub-haplotypes whose p-value for the correlation with the
~~clinical response or outcome of interest is less than or equal to~~ a second selected value,
wherein the second selected value is less than the first selected value.
- 15 68. (currently amended) The method of any one of claims 64-67, further comprising the step
of ~~excluding from further processing discarding a complex redundant sub-haplotype~~
~~subhaplotypes which are~~ is constructed from smaller saved sub-haplotypes, where
wherein the numerical measure of the degree of correlation of each of the smaller saved sub-
haplotypes has correlation values that are is at least as significant as that of the complex
20 redundant sub-haplotype.

69. - 109 (canceled)

110. (currently amended) A computer-usable medium having computer-readable program code
25 stored thereon, for causing a computer to execute a method to determine polymorphic
sites polymorphisms or sub-haplotypes for a locus that correlate with a clinical response or
an outcome of interest, or other phenotype, the computer-readable program code the
method comprising:
(a) ~~computer-readable program code for causing a computer to accessing a database~~
30 containing comprising a haplotype information for each allele of the locus for each
subject in a cohort of subjects, and clinical response or outcome data (clinical outcome

~~values)~~ an outcome value for the outcome of interest for each subject in the or other phenotype data, from a cohort of subjects, the locus comprising at least two polymorphic sites;

- (b) ~~computer-readable program code for causing a computer to statistically analyze~~
5 analyzing each individual SNP-polymorphism in the haplotype-haplotypes for the
degree to which it correlates with the clinical outcome values or other phenotype data,
outcome of interest and generating calculating a numerical measure of the degree of
correlation;
- (c) ~~computer-readable program code for causing a computer to store for further processing~~
10 storing each these individual SNPs-polymorphism whose numerical measure of the
degree of correlation with the clinical outcome values or other phenotype data exceeds
meets a first cut-off value criterion and its numerical measure of the degree of
correlation;
- (d) ~~computer-readable program code for causing a computer to generate-generating all~~
15 possible pair-wise combinations of the saved SNPs-individual polymorphisms so as to
provide a set of n -site sub-haplotypes where $n = 2$;
- (e) ~~computer-readable program code for causing a computer to statistically analyze~~
analyzing each newly generated n -site sub-haplotype for the degree to which it
correlates with the clinical outcome of interest-values or other phenotype data, and
20 calculate-calculating the a numerical measure of the degree of correlation;
- (f) ~~computer-readable program code for causing a computer to store for further processing~~
storing each these n -site sub-haplotypes-haplotype whose numerical measure of the
degree of correlation exceeds-meets the first cut-off value criterion and its numerical
measure of the degree of correlation;
- (g) ~~computer-readable program code for causing a computer to generate-generating all~~
25 possible pair-wise combinations among and between the saved SNPs-individual
polymorphisms and saved sub-haplotypes, to produce new sub-haplotypes with
increased values of n ; and
- (h) ~~computer-readable program code for causing a computer to repeat-repeating steps (e)~~
30 through (g) until either (i) no new sub-haplotypes can be generated, or (ii) no further

new sub-haplotypes having n less than a pre-selected ~~or~~ user-selected limit can be generated.

111. (currently amended) The computer-usable medium of claim 110, wherein the method
5 which further comprises computer-readable program code stored thereon for causing a
computer to displaying display those saved SNPs individual polymorphisms and sub-
haplotypes whose numerical measure of the degree of correlation with the clinical outcome
value or other phenotype exceedsmeets a second cut-off valuecriterion, wherein the second
cut-off value criterion is greater more stringent than the first cut-off valuecriterion.

112. (currently amended) A computer-usable medium having computer-readable program code
stored thereon, for causing a computer to execute a method to determine polymorphic sites
or sub-haplotypes for a locus that correlate with a ~~clinical response or an~~ outcome of
interest, ~~or other phenotype~~, the ~~computer-readable program code~~ method comprising:

15 (a) ~~computer-readable program code for causing a computer to access~~ accessing a database
containing comprising a haplotype for each allele of the locus for each subject in a
cohort of subjects information, and an outcome value for the outcome of interest
clinical response or outcome data (clinical outcome values) for each subject in their
other phenotype data, from a cohort of subjects, the locus comprising at least two
20 polymorphic sites;

(b) ~~computer-readable program code for causing a computer to statistically analyze~~
analyzing each individual SNP polymorphism in the ~~haplotype~~ haplotypes for the
degree to which it correlates with the clinical outcome of interest ~~values or other~~
~~phenotype data, and calculate~~ calculating the p-value for the ~~degree of correlation;~~

25 (c) ~~computer-readable program code for causing a computer to store for further processing~~
storing these each individual SNPs polymorphism whose p-value for the degree of
correlation does not exceed a first cut-off value and its p-value;

(d) ~~computer-readable program code for causing a computer to generate~~ generating all
possible pair-wise combinations of the saved SNPs individual polymorphisms so as to
30 provide a set of n -site sub-haplotypes where $n = 2$;

- (e) ~~computer-readable program code for causing a computer to statistically analyze~~
analyzing each newly generated n -site sub-haplotype for the degree to which it
correlates with the ~~clinical outcome of interest values or other phenotype data, and~~
~~calculate calculating~~ the p-value for the degree of correlation;
- 5 (f) ~~computer-readable program code for causing a computer to store for further processing~~
storing those ~~each~~ n -site sub-haplotypes haplotype whose p-value ~~for the degree of~~
~~correlation~~ does not exceed the first cut-off value and its p-value;
- (g) ~~computer-readable program code for causing a computer to generate~~generating all
possible pair-wise combinations among and between the saved SNPs individual
10 polymorphisms and saved sub-haplotypes, to produce new subhaplotypes with
increased values of n ; and
- (h) ~~computer-readable program code for causing a computer to repeat~~repeating steps (e)
through (g) until either (i) no new sub-haplotypes can be generated, or (ii) no further
new sub-haplotypes having n less than a pre-selected or ~~user-selected~~ limit can be
15 generated.

113. (currently amended) The computer-usable medium of claim 110, ~~which further comprises~~
~~computer-readable program code stored thereon for causing a computer to~~ wherein the
method further comprises displaying ~~display~~ those ~~the~~ saved SNPs individual
20 polymorphisms and sub-haplotypes whose p-value ~~for the degree of correlation with the~~
~~clinical outcome value or other phenotype~~ does not exceed a second cut-off value, wherein
the second cut-off value is less than the first cut-off value.

114. (currently amended) The computer-usable medium of claim~~claims~~ 110-113, ~~which further~~
25 ~~comprises computer-readable program code stored thereon for causing a computer to~~
~~exclude from further processing~~ wherein the method further comprises eliminating a
complex ~~redundant subhaplotypes~~ subhaplotype ~~which are constructed~~generated from
smaller saved sub-haplotypes, wherein the numerical measure of the degree of correlation
of each of ~~where the smaller saved sub-haplotypes each have correlation values that are is~~
30 at least as significant as that of the complex ~~redundant~~ sub-haplotype.

115. (currently amended) A computer-usable medium having computer-readable program code stored thereon, for causing a computer to execute a method to determine polymorphic sites or sub-haplotypes for a locus that correlate with a clinical response or an outcome of interest, or other phenotype of interest, the computer-readable program code method comprising:

(a) ~~computer-readable program code for causing a computer to access~~accessing a database containing single gene comprising a haplotype for each allele of the locus for each subject in a cohort of subjects information for one or more genes, and clinical response, an outcome data, or other phenotype data from a value for the outcome of interest for each subject in the cohort of subjects, the locus comprising at least two polymorphic sites;

(b) ~~computer-readable program code for causing a computer to statistically analyze~~analyzing each single gene haplotype for the degree to which it correlates with the clinical response, outcome, or phenotype of interest, and to generate~~calculating a~~ numerical measure of the degree of correlation;

(c) ~~computer-readable program code for causing a computer to store for further processing~~storing those haplotypes each haplotype whose numerical measure of the degree of correlation exceeds meets a first cut-off value criterion and its numerical measure of the degree of correlation;

(d) ~~computer-readable program code for causing a computer to generate~~generating, for from each haplotype composed of m polymorphic sites, all possible sub-haplotypes having a single site masked, so as to provide a set of $m-n$ site sub-haplotypes where $n = 1$;

(e) ~~computer-readable program code for causing a computer to statistically analyze~~analyzing each newly generated sub-haplotype for the degree to which it correlates with the clinical response, outcome, or phenotype of interest, and calculating a the numerical measure of the degree of correlation;

(f) ~~computer-readable program code for causing a computer to save for further processing~~storing those each sub-haplotypes haplotype whose numerical measure of the degree of correlation exceeds meets the first cut-off value criterion and its numerical measure of

the degree of correlation;

(g) ~~computer-readable program code for causing a computer to generate~~generating, from the ~~saved~~stored sub-haplotypes, all possible sub-haplotypes having one additional site masked; and

5 (h) ~~computer-readable program code for causing a computer to repeat~~repeating steps (e) through (g) until either (i) no new sub-haplotypes ~~have a degree of correlation which exceeds the first cut-off value~~, or (ii) no further sub-haplotypes having more unmasked sites than a pre-selected limit can be generated.

10 116. (currently amended) The computer-usable medium of claim 115, ~~which further comprises computer-readable program code stored thereon for causing a computer to display~~wherein the method further comprises displaying those saved stored haplotypes and sub-haplotypes whose numerical measure of the degree of correlation with the clinical response data, outcome value, or other phenotype data exceeds meets a second cut-off ~~value~~criterion,
15 ~~wherein the second cut-off value criterion is greater~~more stringent than the first cut-off ~~value~~criterion.

117. (currently amended) A computer-usable medium having computer-readable program code stored thereon, for causing a computer to execute a method to determine polymorphic sites
20 or sub-haplotypes for a locus that correlate with a clinical response or an outcome of interest, ~~or other phenotype of interest, the computer-readable program code~~method comprising:

(a) ~~computer-readable program code for causing a computer to access~~accessing a database
25 ~~containing single gene comprising a haplotype for each allele of the locus for each subject in a cohort of subjects information for one or more genes, and clinical response,~~
an outcome value for the outcome of interest for each subject in the data, or other phenotype data from a cohort of subjects, the locus comprising at least two polymorphic site;

30 (b) ~~computer-readable program code for causing a computer to statistically analyze~~
analyzing each single gene haplotype for the degree to which it correlates with the clinical response, outcome, or phenotype of interest, and to calculatecalculating the p-

value for the degree of correlation;

(c) ~~computer-readable program code for causing a computer to store for further processing~~
storing those each haplotypes haplotype whose p-value for the degree of correlation
does not exceed a first cut-off value and its p-value;

5 (d) ~~computer-readable program code for causing a computer to generate~~generating, for
each haplotype composed of m polymorphic sites, all possible sub-haplotypes having a
single site masked, so as to provide a set of $m-n$ site sub-haplotypes where $n = 1$;

(e) ~~computer-readable program code for causing a computer to statistically analyze~~
analyzing each newly generated sub-haplotype for the degree to which it correlates
10 with the clinical response, outcome, or phenotype of interest, and calculating the p-
value for the degree of correlation;

(f) ~~computer-readable program code for causing a computer to save for further processing~~
storing those sub-haplotypes whose p-value for the degree of correlation does not
exceed the first cut-off value and its p-value;

15 (g) ~~computer-readable program code for causing a computer to generate~~generating, from
the ~~saved~~stored sub-haplotypes, all possible sub-haplotypes having one additional site
masked; and

(h) ~~computer-readable program code for causing a computer to repeat~~repeating steps (e)
through (g) until either (i) no new sub-haplotypes have a p-value which does not the
20 first cut-off value, or (ii) no further sub-haplotypes having more unmasked sites than a
pre-selected limit can be generated.

118. (currently amended) The computer-usable medium of claim 117, ~~which wherein the~~
method further comprises computer-readable program code stored thereon for causing a
25 computer to display displaying those saved-stored haplotypes and sub-haplotypes whose p-
value for the degree of correlation with the clinical response, outcome, or phenotype of
interest does not exceed a second cut-off value, wherein the second cut-off value is less
than the first cut-off value.

30 119. (currently amended) The computer-usable medium of claim 117, claims 115-118, which

wherein the method further comprises ~~computer-readable program code stored thereon for~~
~~causing a computer to exclude from further processing discarding a complex redundant~~
~~sub-haplotypes haplotype which are is~~ constructed from smaller stored sub-haplotypes,
wherein the p-value of each of the smaller sub-haplotypes ~~each have correlation values that~~
are is at least as significant as that of the complex redundant sub-haplotype.

120. - 160. (canceled)

161. (currently amended) A computer programmed to execute a method to determine

polymorphic sites or sub-haplotypes for a locus that correlate with a clinical response or an
outcome of interest, ~~or other phenotype~~, the computer comprising a memory having at least
one region for storing computer executable program code and a processor for executing the
program code stored in memory, wherein the ~~program code includes~~ method comprises:

(a) ~~computer-readable program code for causing a computer to access~~ accessing a database
containing comprising a haplotype information, for each allele of the locus for each
subject in a cohort of subjects and an outcome value ~~clinical response or outcome data~~
~~(clinical outcome values) or other phenotype data, from a for the outcome of interest~~
for each subject in the cohort of subjects, the locus comprising at least two
polymorphic sites;

(b) ~~computer-readable program code for causing a computer to statistically analyze~~
analyzing each individual SNP polymorphism in the haplotype for the degree to which
it correlates with the ~~clinical outcome of interest values or other phenotype data;~~ and
generating calculating a numerical measure of the degree of correlation;

(c) ~~computer-readable program code for causing a computer to store for further processing~~
saving these each individual SNPs polymorphism whose numerical measure of the
degree of correlation ~~with the clinical outcome values or other phenotype data~~
exceeds meets a first cut-off ~~value~~ criterion and its numerical measure of the degree of
correlation;

(d) ~~computer-readable program code for causing a computer to generate~~ generating all
possible pair-wise combinations of the saved SNPs individual polymorphisms so as to
provide a set of n -site sub-haplotypes where $n = 2$;

(e) ~~computer-readable program code for causing a computer to statistically analyze~~
analyzing each newly generated n -site sub-haplotype for the degree to which it
correlates with the ~~clinical outcome values or other phenotype data of interest~~ and
~~calculate calculating a the~~ numerical measure of the degree of correlation;

(f) ~~computer-readable program code for causing a computer to store for further processing~~
saving these each n -site sub-haplotypes-haplotype whose numerical measure of the
degree of correlation ~~exceeds-meets~~ the first cut-off ~~value~~criterion and its numerical
measure of the degree of correlation;

(g) ~~computer-readable program code for causing a computer to generate~~generating all
possible pair-wise combinations among and between the saved SNPs individual
polymorphisms and saved sub-haplotypes, to produce new subhaplotypes sub-
haplotypes with increased values of n ; and

(h) ~~computer-readable program code for causing a computer to repeat~~repeating steps (e)
through (g) until either (i) no new sub-haplotypes can be generated, or (ii) no further
new sub-haplotypes having n less than a pre-selected or ~~user-selected~~ limit can be
generated.

162. (currently amended) The computer of claim 161, wherein the ~~program code~~method
further ~~includes-comprises~~ ~~computer-readable program code for causing a computer to~~
displaydisplaying those saved SNPs individual polymorphisms and sub-haplotypes whose
numerical measure of the degree of correlation with the ~~clinical outcome value or other~~
~~phenotype exceeds-meets~~ a second cut-off ~~value~~criterion, wherein the second cut-off ~~value~~
criterion is ~~greater-more stringent~~ than the first cut-off ~~value~~criterion.

163. (currently amended) A computer programmed to execute a method to determine
polymorphic sites or sub-haplotypes for a locus that correlate with a ~~clinical response or an~~
outcome of interest, or ~~other phenotype~~, the computer comprising a memory having at least
one region for storing computer executable program code and a processor for executing the
program code stored in memory, wherein the ~~program code includes~~method comprises:
(a) ~~computer-readable program code for causing a computer to access~~accessing a database
containing-comprising a haplotype for each allele of the locus for each subject in a

cohort of subjects information, and an outcome value for the outcome of interest for each subject in the clinical response or outcome data (clinical outcome values) or other phenotype data, from a cohort of subjects, the locus comprising at least two polymorphic sites;

- 5 (b) ~~computer readable program code for causing a computer to statistically analyze~~
analyzing each individual SNP polymorphism in the haplotype haplotypes for the
degree to which it correlates with the ~~clinical outcome values or other phenotype~~
data of interest and ~~calculate~~ calculating the a p-value for the degree of correlation;
- (c) ~~computer readable program code for causing a computer to store for further processing~~
10 saving these each individual SNPs polymorphism whose p-value for the degree of
correlation does not exceed a first cut-off value and its p-value;
- (d) ~~computer readable program code for causing a computer to generate~~ generating all
possible pair-wise combinations of the saved SNPs individual polymorphisms so as to
provide a set of n -site sub-haplotypes where $n = 2$;
- 15 (e) ~~computer readable program code for causing a computer to statistically analyze~~
analyzing each newly generated n -site sub-haplotype for the degree to which it
correlates with the ~~clinical outcome of interest values or other phenotype data, and~~
calculate calculating the p-value for the degree of correlation;
- (f) ~~computer readable program code for causing a computer to store for further processing~~
20 saving these each n -site sub-haplotypes haplotype whose p-value for the degree of
correlation does not exceed the first cut-off value and its p-value;
- (g) ~~computer readable program code for causing a computer to generate~~ generating all
possible pair-wise combinations among and between the saved SNPs individual
polymorphisms and saved sub-haplotypes, to produce new subhaplotypes sub-
25 haplotypes with increased values of n ; and
- (h) ~~computer readable program code for causing a computer to repeat~~ repeating steps (e)
through (g) until either (i) no new sub-haplotypes can be generated, or (ii) no further
new sub-haplotypes having n less than a pre-selected ~~or user selected~~ limit can be
generated.

164. (currently amended) The computer of claim 161, wherein the ~~program code~~method
further ~~includes comprises~~ computer-readable program code for causing a computer to
~~display~~displaying those saved SNPs individual polymorphisms and sub-haplotypes whose
5 p-value for the degree of correlation with the clinical outcome value or other phenotype
does not exceed a second cut-off value, wherein the second cut-off value is less than the
first cut-off value.

165. (currently amended) The computer of any one of claims 161-~~164~~162, wherein the
10 ~~program code~~ further ~~includes computer-readable program code~~ for causing a computer to
~~exclude from further processing~~ method further comprises discarding a complex redundant
~~subhaplotypes sub-haplotype~~ which are constructed is generated from smaller saved sub-
haplotypes, wherein the numerical measure of the degree of correlation of each of the
smaller saved sub-haplotypes each ~~have correlation values that are~~ is at least as significant
15 as that of the complex redundant sub-haplotype.

166. (currently amended) A computer programmed to execute a method to determine
polymorphic sites or sub-haplotypes for a locus that correlate with a clinical response or an
outcome of interest, ~~or other phenotype of interest~~, the computer comprising a memory
20 having at least one region for storing computer executable program code and a processor
for executing the program code stored in memory, wherein the ~~program code~~
includes method comprises:

(a) ~~computer-readable program code for causing a computer to access~~ accessing a database
containing ~~single gene~~ comprising a haplotype for each allele of the locus for each
25 subject in a cohort of subjects information for one or more genes, and an outcome value
for the outcome of interest for each subject in the ~~clinical response, outcome data, or~~
~~other phenotype data from a cohort of subjects, the locus comprising at least two~~
polymorphic sites;

(b) ~~computer-readable program code for causing a computer to statistically analyze~~
30 analyzing each ~~single gene~~ haplotype for the degree to which it correlates with the
~~clinical response, outcome, or phenotype of interest, and to generate~~ calculating a

numerical measure of the degree of correlation;

(c) ~~computer-readable program code for causing a computer to store for further processing~~
~~saving those each haplotypes~~ haplotype whose numerical measure of the degree of
correlation ~~exceeds~~ meets a first cut-off value criterion and its numerical measure of the
degree of correlation;

(d) ~~computer-readable program code for causing a computer to generate~~ generating, for
from each haplotype composed of m polymorphic sites, all possible sub-haplotypes
having a single site masked, so as to provide a set of $m-n$ site sub-haplotypes where $n =$
1;

(e) ~~computer-readable program code for causing a computer to statistically analyze~~
analyzing each newly generated sub-haplotype for the degree to which it correlates
with the ~~clinical response, outcome, or phenotype~~ of interest, and calculating a the
numerical measure of the degree of correlation;

(f) ~~computer-readable program code for causing a computer to save for further processing~~
~~saving those each sub-haplotypes~~ haplotype whose numerical measure of the degree of
correlation ~~exceeds~~ meets the first cut-off value criterion and its numerical measure of
the degree of correlation;

(g) ~~computer-readable program code for causing a computer to generate~~ generating, from
the saved sub-haplotypes, all possible sub-haplotypes having one additional site
masked; and

(h) ~~computer-readable program code for causing a computer to repeat~~ repeating steps (e)
through (g) until either (i) no new sub-haplotypes ~~have a degree of correlation which~~
~~exceeds the first cut-off value~~, or (ii) no further sub-haplotypes having more unmasked
sites than a pre-selected limit can be generated.

167. (currently amended) The computer of claim 166, wherein the ~~program code~~ method
~~further includes~~ comprises ~~computer-readable program code for causing a computer to~~
~~display~~ displaying those saved haplotypes and sub-haplotypes whose numerical measure of
the degree of correlation with the ~~clinical response data, outcome value, or other phenotype~~
~~data~~ exceeds meets a second cut-off value criterion, wherein the second cut-off value

criterion is greater-more stringent than the first cut-off valuecriterion.

168. (currently amended) A computer programmed to execute a method to determine
polymorphic sites or sub-haplotypes for a locus that correlate with a clinical response or an
outcome of interest, or other phenotype of interest, the computer comprising a memory
having at least one region for storing computer executable program code and a processor
for executing the program code stored in memory, wherein the program code
includesmethod comprises:

(a) ~~computer readable program code for causing a computer to access~~ accessing a database
containing single genecomprising a haplotype for each allele of the locus for each
subject in a cohort of subjectsinformation for one or more genes, and clinical
response,an outcome data, or other phenotype data from a value for the outcome of
interest for each subject in the cohort of subjects, the locus comprising at least two
polymorphic sites;

(b) ~~computer readable program code for causing a computer to statistically analyze~~
analyzing each single gene haplotype for the degree to which it correlates with the
clinical response, outcome, or phenotype of interest, and ~~to calculate~~calculating the a p-
value for the degree of correlation;

(c) ~~computer readable program code for causing a computer to store for further processing~~
saving these each haplotype haplotypes whose p-value for the degree of correlation
does not exceed a first cut-off value and its p-value;

(d) ~~computer readable program code for causing a computer to generate~~generating, for
from each haplotype composed of m polymorphic sites, all possible sub-haplotypes
having a single site masked, so as to provide a set of $m-n$ site sub-haplotypes where $n =$
1;

(e) ~~computer readable program code for causing a computer to statistically analyze~~
analyzing each newly generated sub-haplotype for the degree to which it correlates
with the clinical response, outcome, or phenotype of interest, and calculating the p-
value for the degree of correlation;

(f) ~~computer readable program code for causing a computer to save for further processing~~

~~saving these each sub-haplotypes-haplotype~~ whose p-value for the degree of correlation does not exceed the first cut-off value and its p-value;

(g) ~~computer-readable program code for causing a computer to generate~~generating, from the saved sub-haplotypes, all possible sub-haplotypes having one additional site masked; and

(h) ~~computer-readable program code for causing a computer to repeat~~repeating steps (e) through (g) until either (i) no new sub-haplotypes have a p-value which does not the first cut-off value, or (ii) no further sub-haplotypes having more unmasked sites than a pre-selected limit can be generated.

169. (currently amended) The computer of claim 168, wherein the ~~program code~~method further ~~includes~~comprises computer-readable program code for causing a computer to ~~display~~displaying those saved haplotypes and sub-haplotypes whose p-value for the degree of correlation with the clinical response, outcome, or phenotype of interest does not exceed a second cut-off value, wherein the second cut-off value is less than the first cut-off value.

170. (currently amended) The computer of any one of claims 166-169~~167~~, wherein the ~~program code~~method further includes ~~computer-readable program code for causing a computer to exclude from further processing~~comprises discarding a complex redundant sub-haplotypes-haplotype which ~~are~~is constructed from smaller saved sub-haplotypes, ~~where wherein the numerical measure of the degree of correlation of each of the smaller saved sub-haplotypes each have correlation values that are~~is at least as significant as that of the complex redundant sub-haplotype.

171. - 183. (canceled)

184. (new) The method of claim 59, wherein providing comprises determining the haplotype for each allele of the locus for each subject and acquiring the outcome value for the outcome of interest for each subject.

185. (new) The method of claim 60, further comprising displaying the numerical measure of the degree of correlation of each displayed individual polymorphism or sub-haplotype.

186. (new) The method of claim 64, wherein providing comprises determining the haplotype for each allele of the locus for each subject and acquiring the outcome value for the outcome of interest for each subject.

187. (new) The method of claim 65, further comprising displaying the numerical measure of the degree of correlation of each displayed haplotype or sub-haplotype.

188. (new) A method of determining polymorphic sites or sub-haplotypes for a locus that correlate with an outcome of interest, comprising:

- (a) providing a haplotype for each allele of the locus for each subject in a cohort of subjects and an outcome value for the outcome of interest for each subject in the cohort of subjects, the locus comprising at least two polymorphic sites;
- (b) statistically analyzing each individual polymorphism in the haplotypes for the degree to which it correlates with the outcome of interest and calculating a numerical measure of the degree of correlation;
- (c) saving each individual polymorphism whose numerical measure of the degree of correlation meets a first cut-off criterion for statistical significance;
- (d) generating all possible pair-wise combinations of the saved individual polymorphisms so as to provide a set of n -site sub-haplotypes where $n = 2$;
- (e) statistically analyzing each newly generated n -site sub-haplotype for the degree to which it correlates with the outcome of interest and calculating the numerical measure of the degree of correlation;
- (f) saving each n -site sub-haplotype whose numerical measure of the degree of correlation meets the first cut-off criterion;
- (g) generating all possible pair-wise combinations among and between the saved individual polymorphisms and saved sub-haplotypes, to produce new sub-haplotypes with increased values of n ; and
- (h) repeating steps (e) through (g) until either (i) no new sub-haplotypes or (ii) no new sub-

haplotypes having n less than a pre-selected limit can be generated.

189. (new) The method of claim 188, wherein providing comprises determining the haplotype
for each allele of the locus for each subject and acquiring the outcome value for the
5 outcome of interest for each subject.

190. (new) The method of claim 188, further comprising saving the numerical measure of the
degree of correlation of each saved individual polymorphism or sub-haplotype.

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